

CLAIM AMENDMENTS

This listing of claims will replace all prior versions and listings of claims in the application:

1. (currently amended) A carrier molecule for transporting a polyanionic macromolecule across a membrane of a cell consisting of:
a single ~~biocompatible hydrophilic~~ polyethylene glycol (PEG) backbone polymer;
and
about 4 to about 100 ~~two or more polycationic~~ polyethylenimine (PEI) polymers covalently linked to the ~~biocompatible hydrophilic~~ PEG backbone polymer by linkers.
- 2.– 4. (canceled)
5. (currently amended) The carrier of claim 1, further comprising at least one targeting moiety connected to the ~~biocompatible hydrophilic~~ PEG backbone polymer or to one of the ~~two or more polycationic~~ about 4 to about 100 PEI polymers.
6. (currently amended) The carrier of claim 5, wherein the at least one targeting moiety is selected from the group consisting of a ligand, an antigen, a hapten, biotin, a lectin, galactose, galactosamine, a protein, a histone, a polypeptide, a lipid, a carbohydrate, a vitamin, and a combination thereof.
7. (currently amended) The carrier of claim 1, further comprising at least one lysis agent connected to the ~~biocompatible hydrophilic~~ PEG backbone polymer or to one of the ~~two or more polycationic~~ about 4 to about 100 PEI polymers.
8. (previously presented) The carrier of claim 7, wherein the at least one lysis agent is selected from the group consisting of a viral peptide, a bacterial toxin, a lytic peptide, aleveolysin, bifermentolysin, boutulinolysin, capriciolysin, cereolysin O, chauveolysin, histolyticolysin O, pneumolysin, sealigerolysin, septicolysin O, sordellilysin, streptoslysin O, tenaolysin or thuringolysin O, and active fragments thereof.

9. (previously presented) The carrier of claim 1, wherein the linkers are about 2 to about 100 atoms in length.
10. (previously presented) The carrier of claim 9, wherein the linkers are selected from the group consisting of a hydrocarbon chain, a PEG fragment, a polypeptide, a linear polymer containing an ester bond, a linear polymer containing an amide bond, a linear polymer containing a disulfide bond, a linear polymer containing a hydrozone bond, and a linear polymer containing an oxime bond.
11. (previously presented) The carrier of claim 9, wherein the linkers are about 3 atoms to about 30 atoms in length.
12. (currently amended) The carrier of claim 1, wherein the ~~biocompatible hydrophilic~~ PEG backbone polymer has a molecular weight in the range from about 1,000 to about 1,000,000 daltons and the ~~polycationic~~ about 4 to about 100 PEI polymers have a molecular weight in the range from about 100 to about 100,000 daltons.
13. (currently amended) The carrier of claim 12, wherein the molecular weight of the ~~biocompatible hydrophilic~~ PEG backbone polymer is in the range from about 5,000 to about 100,000 daltons.
14. (currently amended) The carrier of claim 12, wherein the molecular weight of the ~~biocompatible hydrophilic~~ PEG backbone polymer is in the range from about 20,000 to about 40,000 daltons.
15. (currently amended) The carrier of claim 12, wherein the molecular weight of the ~~polycationic~~ about 4 to about 100 PEI polymers is in the range from about 200 to about 10,000 daltons.

16. (currently amended) The carrier of claim 12, wherein the molecular weight of the ~~polycationic~~ about 4 to about 100 PEI polymers is in the range from about 400 to about 2,000 daltons.
17. (canceled)
18. (canceled)
19. (currently amended) The carrier of claim ~~11~~7, wherein from about 8 to about 15 PEI ~~polycationic~~ polymers are covalently linked to the ~~biocompatible~~ PEG backbone polymer by linkers.
20. (currently amended) The carrier of claim ~~11~~7, wherein the molecular weight of the ~~biocompatible hydrophilic~~ PEG backbone polymer is in the range from about 20,000 to about 40,000 daltons.
21. (currently amended) The carrier of claim ~~11~~7, wherein the molecular weight of ~~polycationic~~ the about 4 to about 100 PEI polymers is in the range from about 400 to about 2,000 daltons.
22. (currently amended) The carrier of claim ~~11~~7, wherein the linkers are selected from the group consisting of a hydrocarbon chain, a PEG fragment, a polypeptide, a linear polymer containing an ester bond, a linear polymer containing an amide bond, a linear polymer containing a disulfide bond, a linear polymer containing a hydrozone bond, and a linear polymer containing an oxime bond.
- 23.-26 (canceled)
27. (currently amended) The carrier of claim ~~22~~17, comprising a linker which is a biodegradable peptide.
28. (canceled)

29. (canceled)

30. (currently amended) The carrier of claim ~~2725~~, wherein the biodegradable peptide is selected from the group consisting of GlyPhePheGly (SEQ ID NO.2) and GlyPheLeuGly (SEQ ID NO.1).

31. (currently amended) A complex for transporting a polyanionic macromolecule across a membrane of a cell comprising:

a carrier molecule for delivering the polyanionic macromolecule to the cell, the carrier molecule consisting of a single ~~biocompatible hydrophilic~~ HPMA backbone polymer and two or more ~~polycationic~~ PEI polymers covalently linked to the ~~biocompatible hydrophilic~~ HPMA backbone polymer by linkers; and
a ~~polyanionic macromolecule~~ nucleic acid complexed with the carrier molecule.

32.-35. (canceled)

36. (currently amended) The complex of claim ~~3132~~, wherein the nucleic acid is selected from the group consisting of genomic DNA, plasmid DNA, synthetic DNA, and RNA.

37. (original) The complex of claim ~~3132~~, wherein the nucleic acid is selected from the group consisting of an antisense oligonucleotide, ribozyme, DNase, chimeric RNA/DNA oligonucleotide, phosphorothioate oligonucleotide, 2'-O-methyl oligonucleotide, DNA-PNA conjugate, DNA-morpholino-DNA conjugate, and a combination thereof.

38. (currently amended) The complex of claim 31, wherein the ~~biocompatible hydrophilic~~ HPMA backbone polymer has a molecular weight in the range from about 1,000 daltons to about 1,000,000 daltons and the ~~polycationic~~ PEI polymers have a molecular weight in the range from about 100 daltons to about 100,000 daltons.

39. (currently amended) The complex of claim 38, wherein the molecular weight of the ~~biocompatible-hydrophilic~~ HPMA backbone polymer is in the range from about 20,000 daltons to about 40,000 daltons.
40. (currently amended) ~~the~~ The complex of claim 39, wherein the molecular weight of the ~~polycationic~~ PEI polymers is in the range from about 400 daltons to about 2,000 daltons.
41. (previously presented) The complex of claim 31, wherein the linkers are selected from the group consisting of a hydrocarbon chain, a PEG fragment, a polypeptide, a linear polymer containing an ester bond, a linear polymer containing an amide bond, a linear polymer containing a disulfide bond, a linear polymer containing a hydrozone bond, and a linear polymer containing an oxime bond.
42. (canceled)
43. (canceled)
44. (currently amended) The complex of claim 31, further comprising at least one targeting moiety connected to the ~~biocompatible-hydrophilic~~ HPMA backbone polymer or to one of the two or more ~~polycationic~~ PEI polymers, the at least one targeting moiety selected from the group consisting of a ligand, an antigen, a hapten, biotin, a lectin, galactose, galactosamine, a protein, a histone, a polypeptide, a lipid, a carbohydrate, and a combination thereof.
45. (currently amended) The complex of claim 31, further comprising at least one lysis agent connected to the ~~biocompatible-hydrophilic~~ HPMA backbone polymer or to one of the two or more ~~polycationic~~ PEI polymers, the at least one lysis agent selected from the group consisting of a viral peptide, a bacterial toxin, a lytic peptide, aleveolysin, bifermentolysin, boutulinolysin, capriciolysin, cereolysin O, chauveolysin, histolyticolysin O, pneumolysin, sealigerolysin, septicolysin O, sordellilysin, streptosolysin O, tenaolysin or thuringolysin O, and active fragments thereof.

46. (currently amended) The complex of claim 31, wherein from about 4 to about 100 ~~polycationic~~ PEI polymers are covalently linked to the ~~biocompatible hydrophilic~~ HPMA backbone polymer by linkers.
47. (currently amended) The complex of claim 31, wherein from about 8 to about 15 ~~polycationic~~ PEI polymers are covalently linked to the ~~biocompatible hydrophilic~~ HPMA backbone polymer by linkers.
48. (currently amended) A method of transporting a polyanionic macromolecule across a membrane of a cell comprising:
- (a) complexing the polyanionic macromolecule to a carrier molecule to create a complex, the carrier molecule consisting of a single biocompatible hydrophilic backbone polymer and two or more polycationic polymers covalently linked to the biocompatible hydrophilic backbone polymer by a biodegradable peptide linker which is from about 2 to about 100 atoms in length~~-linkers~~; and
 - (b) contacting the cell with the complex.
49. (original) The method of claim 48, wherein the biocompatible hydrophilic backbone is selected from the group consisting of polyethylene glycol (PEG), poly (N-(2-hydroxypropyl)methacrylamide), and copolymers thereof.
50. (original) The method of claim 49, wherein the polycationic polymers are selected from the group consisting of polyalkylamine (PAM), polyethylenimine (PEI), a polylysine (PL), a polypeptide, chitosan, a polysaccharide, and copolymers thereof.
51. (currently amended) The method of claim 48, further comprising at least one targeting moiety connected to the biocompatible hydrophilic backbone or to one of the two or more polycationic polymers, the at least one targeting moiety selected from the group consisting of a ligand, an antigen, a hapten, biotin, a lectin, galactose, galactosamine, a protein, a histone, a polypeptide, a lipid, a carbohydrate, and a combination thereof.

52. (original) The method of claim 48, further comprising at least one lysis agent connected to the biocompatible hydrophilic backbone or to one of the two or more polycationic polymers, the at least one lysis agent selected from the group consisting of a viral peptide, a bacterial toxin, a lytic peptide, aleveolysin, bifermentolysin, boutulinolysin, capriciolysin, cereolysin O, chauveolysin, histolyticolysin O, pneumolysin, sealigerolysin, septicolysin O, sordellilysin, streptoslysin O, tenaolysin or thuringolysin O, and active fragments thereof.
53. (canceled)
54. (canceled)
55. (canceled)
56. (currently amended) The method of claim ~~48~~55, wherein the biodegradable peptide linker is selected from the group consisting of GlyPhePheGly (SEQ ID NO.2) and GlyPheLeuGly (SEQ ID NO.1).
57. (previously presented) The method of claim 48, wherein the biocompatible hydrophilic backbone has a molecular weight in the range from about 1,000 to about 1,000,000 daltons and the polycationic polymers have a molecular weight in the range from about 100 to about 100,000 daltons.
58. (previously presented) The method of claim 57, wherein the molecular weight of the biocompatible hydrophilic backbone is in the range from about 20,000 to about 40,000 daltons.
59. (previously presented) The method of claim 57, wherein the molecular weight of the polycationic polymers is in the range from about 400 to about 2,000 daltons.

60. (original) The method of claim 57, wherein the biocompatible hydrophilic backbone is polyethylene glycol and the polycationic polymers are polyethylenimine.
61. (previously presented) The method of claim 60, wherein the molecular weight of the biocompatible hydrophilic backbone is in the range from about 20,000 to about 40,000 daltons.
62. (previously presented) The method of claim 60, wherein the molecular weight of the polycationic polymers is in the range from about 400 to about 2,000 daltons.
63. (currently amended) The method of claim 48, wherein from about 4 to about 100 polycationic polymers are covalently linked to the biocompatible hydrophilic backbone polymer by the biodegradable peptide linker~~linkers~~.
64. (currently amended) The method of claim 48, wherein from about 8 to about 15 polycationic polymers are covalently linked to the biocompatible hydrophilic backbone polymer by the biodegradable peptide linker~~linkers~~.
- 65-83. canceled
84. (currently amended) A carrier for transporting a polyanionic macromolecule across a membrane of a cell comprising:
a PEG biocompatible hydrophilic backbone polymer; and
two or more PEI polycationic polymers covalently linked to the PEG biocompatible hydrophilic backbone polymer by peptide or propionic acid linkers;
wherein the PEI polycationic polymers have a molecular weight of from about 400 to about 2,000 daltons.
85. (previously presented) The carrier of claim 84 wherein the PEI polycationic polymers have a molecular weight of about 2,000 daltons.

86. (previously presented) The carrier of claim 84 wherein the PEI polycationic polymers have a molecular weight of about 1,200 daltons.

87. (previously presented) The carrier of claim 84 wherein the PEI polycationic polymers have a molecular weight of about 800 daltons.

88. (previously presented) The carrier of claim 84 wherein the PEI polycationic polymers have a molecular weight of about 400 daltons.

89.-94. (canceled)

95. (new) The carrier of claim 8494, wherein the linker is a peptide and the peptide is selected from the group consisting of GlyPhePheGly (SEQ ID NO. 2) and GlyPheLeuGly (SEQ ID NO. 1).

96. (previously presented) The carrier of claim 84, wherein the molecular weight of the PEG biocompatible hydrophilic backbone polymer is in the range from about 20,000 to about 40,000 daltons.